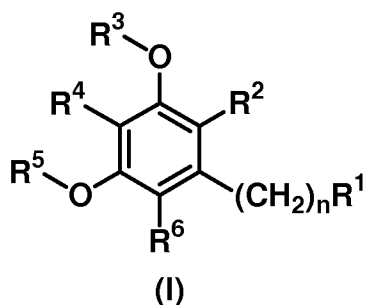


a.) Amendment to the Claims

1. (Currently Amended and withdrawn) ~~An Hsp90 family protein inhibitor comprising, as an active ingredient,~~ A method of inhibiting a heat shock protein 90 family protein, which comprises administering to a patient, in need thereof, an effective amount of a benzene derivative represented by formula (I):



{ wherein

n represents an integer of 0 to 10;

R¹ represents a hydrogen atom, a hydroxy, a cyano, a carboxy, a nitro, a halogen, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkynyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkoxy carbonyl, a substituted or unsubstituted aroyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted heterocyclic-alkyl, a substituted or unsubstituted aryl, a substituted or unsubstituted aralkyl, a substituted or unsubstituted arylsulfonyl, a substituted or unsubstituted heterocyclic group, -CONR⁷R⁸ (wherein R⁷ and R⁸, which may be the same or different, each represent a hydrogen atom, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkanoyl, a

substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group, a substituted or unsubstituted aralkyl a substituted or unsubstituted heterocyclic-alkyl or a substituted or unsubstituted aroyl, or R⁷ and R⁸ form a substituted or unsubstituted heterocyclic group together with the adjacent nitrogen atom), -NR⁹R¹⁰ [wherein R⁹ and R¹⁰, which may be the same or different, each represent a hydrogen atom, a substituted or unsubstituted lower alkylsulfonyl, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group, a substituted or unsubstituted aralkyl, a substituted or unsubstituted heterocyclic-alkyl, a substituted or unsubstituted aroyl, or -CONR¹¹R¹² (wherein R¹¹ and R¹² have the same meanings as the above R⁷ and R⁸, respectively), or R⁹ and R¹⁰ form a substituted or unsubstituted heterocyclic group together with the adjacent nitrogen atom], or -OR¹³ (wherein R¹³ represents a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group, a substituted or unsubstituted aralkyl or a substituted or unsubstituted heterocyclic-alkyl);

R² represents a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted aryl or a substituted or unsubstituted heterocyclic group (but excepting a substituted or unsubstituted pyrazolyl);

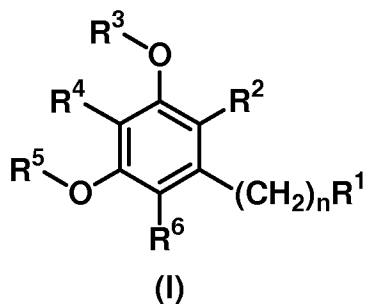
R³ and R⁵, which may be the same or different, each represent a hydrogen atom, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkylsulfonyl, a substituted or

unsubstituted arylsulfonyl, a carbamoyl, a sulfamoyl, a substituted or unsubstituted lower alkylaminocarbonyl, a substituted or unsubstituted di-lower alkylaminocarbonyl, a substituted or unsubstituted lower alkoxy carbonyl, a substituted or unsubstituted heterocyclic-carbonyl, a substituted or unsubstituted aralkyl or a substituted or unsubstituted aroyl; and

R^4 and R^6 , which may be the same or different, each represent a hydrogen atom, a hydroxy, a halogen, a cyano, a nitro, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkynyl, a substituted or unsubstituted lower alkoxy, a substituted or unsubstituted cycloalkyl, an amino, a lower alkylamino, a di-lower alkylamino, a carboxy, a substituted or unsubstituted lower alkoxy carbonyl, a substituted or unsubstituted aryloxy, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group (but excepting a substituted or unsubstituted pyrazolyl), a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aralkyl, or a substituted or unsubstituted heterocyclic-alkyl}, or

a prodrug thereof, or a pharmaceutically acceptable salt thereof.

2. (Currently Amended and withdrawn) ~~An Hsp90 family protein inhibitor comprising, as an active ingredient,~~ A method of inhibiting a heat shock protein 90 family protein, which comprises administering to a patient, in need thereof, an effective amount of a benzene derivative represented by general formula (I):



(wherein

n represents an integer of 0 to 10;

R^1 represents a hydrogen atom, a hydroxy, a cyano, a carboxy, a nitro, a halogen, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkynyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkoxy carbonyl, a substituted or unsubstituted aroyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted heterocyclic-alkyl, a substituted or unsubstituted aryl, a substituted or unsubstituted aralkyl, a substituted or unsubstituted arylsulfonyl, a substituted or unsubstituted heterocyclic group, $-\text{CONR}^7\text{R}^8$ (wherein R^7 and R^8 , which may be the same or different, each represent a hydrogen atom, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group, a substituted or unsubstituted aralkyl, a substituted or unsubstituted heterocyclic-alkyl or a substituted or unsubstituted aroyl, or R^7 and R^8 form a substituted or unsubstituted heterocyclic group together with the adjacent nitrogen atom), $-\text{NR}^9\text{R}^{10}$ [wherein R^9 and R^{10} , which may be the same or different, each represent a hydrogen atom, a substituted or

unsubstituted lower alkylsulfonyl, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group, a substituted or unsubstituted aralkyl, a substituted or unsubstituted heterocyclic-alkyl, a substituted or unsubstituted aroyl, or $-\text{CONR}^{11}\text{R}^{12}$ (wherein R^{11} and R^{12} have the same meanings as the above R^7 and R^8 , respectively), or R^9 and R^{10} form a substituted or unsubstituted heterocyclic group together with the adjacent nitrogen atom], or $-\text{OR}^{13}$ (wherein R^{13} represents a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group, a substituted or unsubstituted aralkyl or a substituted or unsubstituted heterocyclic-alkyl);

R^2 represents a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted aryl or a substituted or unsubstituted heterocyclic group (but excepting a substituted or unsubstituted pyrazolyl);

R^3 and R^5 , which may be the same or different, each represent a hydrogen atom, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkylsulfonyl, a substituted or unsubstituted arylsulfonyl, a carbamoyl, a sulfamoyl, a substituted or unsubstituted lower alkylaminocarbonyl, a substituted or unsubstituted di-lower alkylaminocarbonyl, a substituted or unsubstituted lower alkoxycarbonyl, a substituted or unsubstituted heterocyclic-carbonyl, a substituted or unsubstituted aralkyl or a substituted or unsubstituted aroyl; and

R⁴ and R⁶, which may be the same or different, each represent a hydrogen atom, a hydroxy, a halogen, a cyano, a nitro, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkynyl, a substituted or unsubstituted lower alkoxy, a substituted or unsubstituted cycloalkyl, an amino, a lower alkylamino, a di-lower alkylamino, a carboxy, a substituted or unsubstituted lower alkoxycarbonyl, a substituted or unsubstituted aryloxy, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group (but excepting a substituted or unsubstituted pyrazolyl), a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aralkyl, or a substituted or unsubstituted heterocyclic-alkyl)) or a pharmaceutically acceptable salt thereof.

3. (Currently Amended and withdrawn) ~~The Hsp90 family protein inhibitor according to claim 2~~ The method according to claim 2, wherein R¹ is a hydrogen atom, a hydroxy, a cyano, a carboxy, a nitro, a halogen, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkynyl, a substituted or unsubstituted lower alkoxy, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkoxycarbonyl, a substituted or unsubstituted lower alkanoyloxy, a substituted or unsubstituted heterocyclic-alkyl, a substituted or unsubstituted aryl, a substituted or unsubstituted arylsulfonyl, -CONR⁷R⁸ or -NR⁹R¹⁰.

4. (Currently Amended and withdrawn) ~~The Hsp90 family protein inhibitor according to claim 2~~ The method according to claim 2, wherein R¹ is a substituted

or unsubstituted lower alkyl, a substituted or unsubstituted lower alkynyl, a substituted or unsubstituted lower alkoxy, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkoxy carbonyl, a substituted or unsubstituted heterocyclic-alkyl, a substituted or unsubstituted aryl, -CONR⁷R⁸, or -NR⁹R¹⁰.

5. (Currently Amended and withdrawn) ~~The Hsp90 family protein inhibitor according to claim 3 or 4~~ The method according to claim 3 or 4, wherein R² is a substituted or unsubstituted aryl, or a substituted or unsubstituted aromatic heterocyclic group.

6. (Currently Amended and withdrawn) ~~The Hsp90 family protein inhibitor according to claim 3 or 4~~ The method according to claim 3 or 4, wherein R² is a substituted or unsubstituted aryl.

7. (Currently Amended and withdrawn) ~~The Hsp90 family protein inhibitor according to claim 3 or 4~~ The method according to claim 3 or 4, wherein R² is a substituted or unsubstituted phenyl.

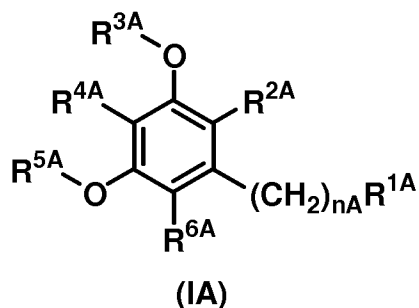
8. (Currently Amended and withdrawn) ~~The Hsp90 family protein inhibitor according to claim 3 or 4~~ The method according to claim 3 or 4, wherein R² is a substituted or unsubstituted furyl.

9. (Currently Amended and withdrawn) ~~The Hsp90 family protein inhibitor according to claim 1 or 2~~ The method according to claim 1 or 2, wherein R⁴ is a hydrogen atom, a hydroxy, or a halogen.

10. (Currently Amended and withdrawn) ~~The Hsp90 family protein inhibitor according to claim 1 or 2~~ The method according to claim 1 or 2, wherein R³ and R⁵, which may be the same or different, each are a hydrogen atom, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aroyl, a substituted or unsubstituted lower alkylaminocarbonyl, a substituted or unsubstituted di-lower alkylaminocarbonyl, a substituted or unsubstituted lower alkoxycarbonyl, or a substituted or unsubstituted heterocyclic-carbonyl.

11. (Currently Amended and withdrawn) ~~The Hsp90 family protein inhibitor according to claim 1 or 2~~ The method according claim 1 or 2, wherein R³, R⁴ and R⁵ are hydrogen atoms.

12. (Currently Amended) A benzene derivative represented by general formula (IA):



[wherein R^{2A} represents ~~a substituted or unsubstituted aryl, or a substituted or unsubstituted aromatic heterocyclic group (but excepting a substituted or unsubstituted pyrazolyl)~~ a substituted or unsubstituted phenyl];

R^{3A} and R^{5A}, which may be the same or different, each represent a hydrogen atom, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkanoyl, a carbamoyl, a sulfamoyl, a substituted or unsubstituted lower alkylsulfonyl, a substituted or unsubstituted lower alkylaminocarbonyl, a substituted or unsubstituted di-lower alkylaminocarbonyl, a substituted or unsubstituted lower alkoxy carbonyl, a substituted or unsubstituted heterocyclic-carbonyl, a substituted or unsubstituted aralkyl, or a substituted or unsubstituted aroyl;

R^{4A} represents a hydrogen atom, a hydroxy, or a halogen;

nA represents an integer of 0 to 5;

provided that;

(1) when n_A is 0,

then R^{1A} is a hydrogen atom, a methyl, a hydroxy, a methoxy, a carboxyl, a methoxycarbonyl, a carbamoyl, $-\text{CONHCH}_3$, $-\text{CON}(\text{CH}_3)_2$, $-\text{CONHCH}_2\text{Ph}$ (wherein Ph represents a phenyl), $-\text{CH}(\text{OCH}_3)\text{Ph}$ (wherein Ph has the same meaning as that defined above), a propionyl, a benzoyl, a dioxolanyl, a substituted or unsubstituted vinyl, or a substituted or unsubstituted prop-1-en-1-yl;

and when R^{1A} is a hydrogen atom,

then R^{6A} is a substituted or unsubstituted lower alkyl;

when R^{1A} is a methyl, a hydroxy, a methoxy, a carboxyl, a methoxycarbonyl, a carbamoyl, $-\text{CONHCH}_3$, $-\text{CON}(\text{CH}_3)_2$, $-\text{CONHCH}_2\text{Ph}$ (wherein Ph has the same meaning as that defined above), a propionyl, a benzoyl, a dioxolanyl, a substituted or unsubstituted vinyl, or a substituted or unsubstituted prop-1-en-1-yl,

then R^{6A} is a halogen;

(2) when n_A is an integer of 1 to 5,

then R^{1A} is a hydroxy, a cyano, a carboxyl, a halogen, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkynyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted lower alkoxycarbonyl, a substituted or unsubstituted aryl, a substituted or unsubstituted aroyl, a substituted or

unsubstituted heterocyclic-alkyl, a substituted or unsubstituted aralkyl, a substituted or unsubstituted arylsulfonyl, a substituted or unsubstituted heterocyclic group, $-\text{CONR}^7\text{R}^8$ (wherein R^7 and R^8 , which may be the same or different, each represent a hydrogen atom, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group, a substituted or unsubstituted aralkyl a substituted or unsubstituted heterocyclic-alkyl or a substituted or unsubstituted aroyl, or R^7 and R^8 form a substituted or unsubstituted heterocyclic group together with the adjacent nitrogen atom), $-\text{NR}^9\text{R}^{10}$ (wherein R^9 and R^{10} , which may be the same or different, each represent a hydrogen atom, a substituted or unsubstituted lower alkylsulfonyl, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group, a substituted or unsubstituted aralkyl, a substituted or unsubstituted heterocyclic-alkyl, a substituted or unsubstituted aroyl), or $-\text{OR}^{13}$ (wherein R^{13} represents a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group, a substituted or unsubstituted aralkyl or a substituted or unsubstituted heterocyclic-alkyl), R^{6A} is a hydrogen atom, a halogen, a cyano, a nitro, a substituted or unsubstituted lower alkyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkynyl, a substituted or unsubstituted lower alkoxy, a substituted or unsubstituted cycloalkyl, a substituted or unsubstituted lower alkanoyl, an amino, a lower alkylamino, a di-lower alkylamino, a carboxy, a substituted or unsubstituted lower alkoxycarbonyl, a substituted or

unsubstituted aryloxy, a substituted or unsubstituted aryl, a substituted or unsubstituted heterocyclic group (but excepting a substituted or unsubstituted pyrazolyl), a substituted or unsubstituted aralkyl, or a substituted or unsubstituted heterocyclic-alkyl;

and provided that;

(i) when R^{3A} and R^{5A} are isopropyl,

then R^{6A} is not a hydrogen atom;

(ii) when R^{3A} and R^{5A} are methyl,

then R^{6A} is not a group selected from a hydrogen atom, a bromo, an ethyl, a 1-hydroxyethyl, a 1-(dimethylamino)ethyl, a vinyl and a carboxy;

(iii) when R^{4A} and R^{6A} are hydrogen atoms, and when R^{3A} and R^{5A} are the same and are tert-butyl or benzyl,

then $-(CH_2)_nR^{1A}$ is not a group selected from a hydroxymethyl and a 2-chloroallyl;

(iv) when R^{4A} and R^{6A} are hydrogen atoms, and when R^{3A} is a benzyl or an acetyl and R^{5A} is a methyl,

or when R^{3A} , R^{4A} and R^{6A} are hydrogen atoms, and when R^{5A} is a methyl,

then $-(CH_2)_nR^{1A}$ is not a group selected from a 2-(acetilamino)propyl and a 2-(substituted lower alkanoylamino)propyl;

(v) when R^{3A} , R^{4A} and R^{5A} are hydrogen atoms, and when R^{6A} is a carboxy, or when R^{4A} , R^{5A} and R^{6A} are hydrogen atoms, and when R^{3A} is a methyl,

then $-(CH_2)_{nA}R^{1A}$ is not an n-pentyl;

(vi) when R^{3A} and R^{4A} are hydrogen atoms, R^{5A} is a methyl, and R^{6A} is an ethyl,

then $-(CH_2)_{nA}R^{1A}$ is not an n-propyl;

(vii) when R^{3A} is a methyl, R^{4A} and R^{6A} are hydrogen atoms, and R^{5A} is a 4-methoxybenzyl,

then $-(CH_2)_{nA}R^{1A}$ is not a group selected from $-(CH_2)_3CH=CH_2$ and $-(CH_2)_5CH=CH_2$;

(viii) when R^{3A} , R^{4A} , R^{5A} and R^{6A} are hydrogen atoms, and when $-(CH_2)_{nA}R^{1A}$ is

(a) an n-pentyl,

then R^{2A} is not a 2,4-dihydroxy-6-pentylphenyl,

~~(b) an n-hexyl,~~

~~then R^{2A} is not a group selected from a 4,6-di(substituted phenyl)triazol-2-yl and a 3,6-di(substituted phenyl)-1,2,4-triazin-5-yl,~~

~~(c) an n-heptyl,~~

~~then R^{2A} is not a substituted triazolyl;~~

~~(ix) when R^{3A} is a hydrogen atom or an acetyl, R^{5A} is a methyl, and R^{4A} and R^{6A} are hydrogen atoms, and when (CH₂)_{nA}R^{1A} is an ethyl or an n-propyl,~~

~~then R^{2A} is not a 2-aminopyrimidin-4-yl having a substituent at the 5-position thereof,~~

~~(x) when R^{3A}, R^{4A} and R^{5A} are hydrogen atoms, R^{6A} is a methoxy, and (CH₂)_{nA}R^{1A} is a 3-methylbut-2-en-1-yl, or a 3-hydroxy-3-methylbutyl,~~

~~then R^{2A} is not a group selected from a 7-hydroxy-4-oxo-4H-1-benzopyran-3-yl and a 6-methoxy-2,2-dimethyl-2H-1-benzopyran-8-yl],~~

or a pharmaceutically acceptable salt thereof.

13. (Currently Amended) The benzene derivative according to claim 12, wherein R^{2A} is a substituted ~~or unsubstituted~~ phenyl, or a pharmaceutically acceptable salt thereof.

14. (Currently Amended) The benzene derivative according to claim 12, wherein R^{2A} is ~~a substituted or unsubstituted furyl~~ phenyl, or a pharmaceutically acceptable salt thereof.

15. (Original) The benzene derivative according to any of claims 12 to 14, wherein R^{3A} and R^{5A} , which may be the same or different, each are a hydrogen atom, a substituted or unsubstituted lower alkanoyl, a substituted or unsubstituted aroyl, a substituted or unsubstituted lower alkenyl, a substituted or unsubstituted lower alkylaminocarbonyl, a substituted or unsubstituted di-lower alkylaminocarbonyl, a substituted or unsubstituted lower alkoxy carbonyl, or a substituted or unsubstituted heterocyclic-carbonyl, or a pharmaceutically acceptable salt thereof.

16. (Original) The benzene derivative according to any of claims 12 to 14, wherein R^{3A} , R^{4A} and R^{5A} are hydrogen atoms, or a pharmaceutically acceptable salt thereof.

17. (Original) The benzene derivative according to any of claims 12 to 14, wherein nA is an integer of 1 to 5, or a pharmaceutically acceptable salt thereof.

18. (Previously Presented) A pharmaceutical composition comprising, as an active ingredient, the benzene derivative according to any of claims 12 to 14 or a pharmaceutically acceptable salt thereof together with a pharmaceutically acceptable carrier.

Claims 19-26 (Cancelled).

27. (Withdrawn and Currently Amended) A method of inhibiting a heat shock protein 90 family protein, which comprises administering ~~an effective amount of a~~ said benzene derivative according to any one of claims 1-4, ~~or a prodrug or a pharmaceutically acceptable salt thereof, to a patient in need thereof~~ or 12-14.

Claims 28-41 (Cancelled).

42. (New) A method of inhibiting a heat shock protein 90 family protein, which comprises administering said prodrug according to claim 1.

43. (New) A method of inhibiting a heat shock protein 90 family protein, which comprises administering said pharmaceutically acceptable salt according to any one of claims 1-4 or 12-14.